✓ATENT COOPERATION TREATY

	From the INTERNATIONAL BUREAU
PCT	То:

NOTIFICATION OF ELECTION

(PCT Rule 61.2)

United States Patent and Trademark Office (Box PCT) Crystal Plaza 2 Washington, DC 20231 ÉTATS-UNIS D'AMÉRIQUE

	ETATS-UNIS D'AMERIQUE
Date of mailing (day/month/year) 09 July 1999 (09.07.99)	in its capacity as elected Office
International application No. PCT/US98/25791	Applicant's or agent's file reference LJIEM1100WO
International filing date (day/month/year) 04 December 1998 (04.12.98)	Priority date (day/month/year) 05 December 1997 (05.12.97)
Applicant	
BOURDON, Mario, A. et al	<u> </u>

1.	The designated Office is hereby notified of its election made:
	X in the demand filed with the International Preliminary Examining Authority on:
	04 June 1999 (04.06.99)
	in a notice effecting later election filed with the International Bureau on:
2.	The election X was
	was not
	made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer D. Barmes	
Facsimile No.: (41-22) 740.14.35	Telephone No.: (41-22) 338.83.38	

PATENT COOPERATION TREATY

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INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

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Applicant's or agent's file reference	FOR EURTHER ACTION	See Notificat	ion of Tonomittal of International		
LJIEM1100WO	FOR FURTHER ACTION	CTION See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416)			
International application No.	International filing date (day/m	onth/year) P	riority date (day/month/year,		
PCT/US98/25791	04 DECEMBER 1998		05 DECEMBER 1997		
International Patent Classification (IPC) of Please See Supplemental Sheet.	or national classification and IPC				
Applicant LA JOLLA INSTITUTE FOR EXPERII	MENTAL MEDICINE				
 This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36. This REPORT consists of a total of sheets. This report is also accompanied by ANNEXES, i.e., sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority. 					
These annexes consist of a to	ion 607 of the Administrative I tal of sheets.	nstructions unde	er the PC1).		
3. This report contains indication	s relating to the following ite	ems:			
I X Basis of the repor					
II Priority	•				
<u></u>					
		elty, inventive	step or industrial applicability		
IV Lack of unity of i					
V X Reasoned statement citations and explan	t under Article 35(2) with regal nations supporting such stateme	rd to novelty, in ent	ventive step or industrial applicability;		
VI Certain documents of	eited				
VII Certain defects in th	e international application	•			
VIII Certain observations	on the international application	n			
Date of submission of the demand Date of completion of this report					
or submission of the demand	Date of	of completion of	this report		
04 JUNE 1999	27	FEBRUARY 20	00		
Name and mailing address of the IPEA/US Authorized officer JOYCE BRIDGERS					
Commissioner of Patents and Tradema Box PCT			PARALEGAL SPECIALIST		
Washington, D.C. 20231	SU	SAN UNGAR	CHEMICAL MATRIX		
Facsimile No. (703) 305-3230	Teleph	one No. (703)	308-0196 / JOB Cm		

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.
PCT/US98/25791

I. Basis of the report					
1. This report has been drawn on the basis of (Substitute sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to the report since they do not contain amendments):					
	X		al application as origina	• •	
	X	the description	, pages <u>1-18</u>	, as originally filed.	
		-		_ , filed with the demand.	
				_ , filed with the letter of	
				, filed with the letter of	
	X	the claims,	Nos. <u>1-29</u>	, as originally filed.	
			Nos. NONE	, as amended under Article 19.	
			Nos. NONE	, filed with the demand.	
			Nos. NONE	, filed with the letter of	
			Nos	, filed with the letter of	
	x	the drawings,	sheets /fig 1-3	, as originally filed.	
	_			, filed with the demand.	
			sheets/ fig NONE	, filed with the letter of	
				, filed with the letter of	
	X X	the description, the claims, the drawings,	Nos. none sheets/fig none		
3.	This to go	report has been es beyond the disclo	stablished as if (some of) sure as filed, as indicated	the amendments had not been made, since they have been considered in the Supplemental Box Additional observations below (Rule 70.2(c)).	
4. Add		l observations, if	necessary:		

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

PCT/US98/25791

V.	. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial ap	pplicability:
	citations and explanations supporting such statement	,

Claims	13-20, 23-25, 27-29	YES
Claims	1-12, 21-22, 26	NO
Claims	13-20, 23-25, 27-29	YES
Claims	1-12, 21-22, 26	NO NO
Claims	1-29	YES
Claims	NONE	NO
	Claims Claims Claims	Claims 1-12, 21-22, 26 Claims 13-20, 23-25, 27-29 Claims 1-12, 21-22, 26 Claims 1-29

2. CITATIONS AND EXPLANATIONS

Claims 1-2 lack novelty under PCT Article 33(2) as being anticipated by Borgstrom (1993). Borgstrom ('93) discloses a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of an agent that inhibits angiogenesis (Linomide) to a cellular system is interpreted to meet the claim limitation "inhibiting a host cell angiogenic effect."

Claims 1-5 and 26 lack novelty under PCT Article 33(2) as being anticipated by either of Ferrara or Borgstrom. Both references disclose a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of an agent that inhibits angiogenesis (anti-VEGF) is interpreted to meet the claim limitation "inhibiting a host cell angiogenic effect."

Claims 1-4 lack novelty under PCT Article 33(2) as being anticipated by either of DiPetro or Hu. Both references disclose a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of an agent that inhibits angiogenesis (TSP1, as in DiPetro, or IL-1 receptor antagonist, antiserum to IL-8, anti-TNF alpha, or monoclonal bFGF, as in Hu)is interpreted to meet the claim limitation "inhibiting a host cell angiogenic effect."

Claims 1-12, 21-22 and 26 lack novelty under PCT Article 33(2) as being anticipated by Lokeshwar. Lokeshwar discloses a method that is inherently the same as that claimed, a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of the 5A1 monoclonal antibody to M-CSF-1, which inherently "inhibits a host cell angiogenic effect."

Claims 1-29 have industrial applicability as defined by PCT Article 33(4).

Claims 13-20, 23-25 and 27-29 meet the criteria set out in PC1 (Continued on Supplemental Sheet.)

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

THE RECIPIENT OF THE PROPERTY	PCT/US98/25791
Supplemental Box (To be used when the space in any of the preceding boxes is not sufficient)	
Continuation of: Boxes I - VIII	Sheet 10
CLASSIFICATION: The International Patent Classification (IPC) and/or the National classifical IPC(7): A61K 39/395, 48/00; C07K 16/18, 16/22, 16/24, 16/26, 16/28 and US CI 388.85	tion are as listed below:: 424/130.1; 514/44; 530/387.1, 388.23,
V. 2. REASONED STATEMENTS - CITATIONS AND EXPLANATIONS (Continuanticle 33(2)-(3), because the prior art does not teach or fairly suggest the claimed in the state of the state	nued): method of inhibiting angiogenesis.
NEW CITATIONS ————————————————————————————————————	bodies to Murine Macrophage Colony- ges 483-488, see entire document.
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INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/25791

A. CLASSIFICATION OF SUBJECT MATTER IPC(6) :A61K 39/395, 48/00; C07K 16/18, 16/22, 16/24, 16/26, 16/28 US CL :424/130.1; 514/44; 530/387.1, 388.23, 388.85 According to International Patent Classification (IPC) or to both national classification and IPC				
	LDS SEARCHED		-	
Minimum d	locumentation searched (classification system followe	d by classification symbols)		
	424/130.1; 514/44; 530/387.1, 388.23, 388.85			
Documenta	tion searched other than minimum documentation to the	e extent that such documents are included	in the fields searched	
Electronic d	data base consulted during the international search (na	ame of data base and, where practicable	, search terms used)	
	N, REGISTRY, CAPLUS, MEDLINE, EMBASE, LIF of angiogenesis, macrophages, M-CSF, M-CSF-1	ESCI, BIOTECHDS, WIPIDS		
C. DOC	CUMENTS CONSIDERED TO BE RELEVANT			
Category*	Citation of document, with indication, where ap	propriate, of the relevant passages	Relevant to claim No.	
X	Database PREV, 1993: 44131748, Inhibition of Angiogensis by the Quinol FASEB Journal. 28 March-April 1993, see entire document.	ine-2-carboxamide Linomide.	1-3, 13-14, 26	
X	FERRARA, Napoleone. The Role of Vascular Endothelial Growth Factor in Pathological Angiogenesis. Breast Cancer Research and Treatment. 1995, Vol. 36, pages 127-137, see entire document.			
х	DiPIETRO, L. et al. Angiogenic Angiogenic Inhibitor Thrombospondin Pathology. September 1993, Vol. 143 entire document.	n 1. American Journal of	!-4, 26	
X Furth	her documents are listed in the continuation of Box C	. See patent family annex.		
'A' do	pecial categories of cited documents: Document defining the general state of the art which is not considered be of particular relevance	"T" later document published after the inte date and not in conflict with the app the principle or theory underlying the	lication but cited to understand	
	oe or particular relevance	*X* document of particular relevance; th	e claimed invention cannot be	
cit	"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other			
special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "O" document referring to an oral disclosure, use, exhibition or other means "O" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art				
"P" document published prior to the international filing date but later than "&" document member of the same patent family the priority date claimed				
	Date of the actual completion of the international search 23 MARCH 1999 Date of mailing of the international search report 15 APR 1999			
Commission Box PCT	mailing address of the ISA/US oner of Patents and Trademarks on, D.C. 20231	Authorized officer NANCYA: JOHNSON	1	
Facsimile 1		Telephone No. (703) 308-0196	you	

INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/25791

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
x	BORGSTROM, P. et al. Complete Inhibition of Angiogenesis and Growth of Microtumors by Anti-Vascular Endothelial Growth Factor Neutralizing Antibody: Novel Concepts of Angiostatic Therapy from Intravital Videomicroscopy. Cancer Research. 01 September 1996, Vol. 56, pages 4032-4039, see entire document.	1-5, 26
X	HU, D.E., et al. Inhibition of Angiogenesis in Rats by IL-1 Receptor Antagonist and Selected Cytokine Antibodies. Inflammation. 1994, Vol. 18, No.1, pages 45-58, see entire doi:iumnet.	1-4